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NEWS	1		0.5	Web Page for STN Seminar Schedule - N. America
NEWS		AUG		CAS REGISTRY enhanced with new experimental property tags
NEWS		AUG		FSTA enhanced with new thesaurus edition
NEWS	4	AUG	13	CA/CAplus enhanced with additional kind codes for granted patents
NEWS	5	AUG	20	CA/CAplus enhanced with CAS indexing in pre-1907 records
NEWS	6	AUG		Full-text patent databases enhanced with predefined
112110		1100		patent family display formats from INPADOCDB
NEWS	7	AUG	27	USPATOLD now available on STN
NEWS	8	AUG		CAS REGISTRY enhanced with additional experimental
MEMO	0	MUG	20	spectral property data
NEWS	9	SEP	0.7	STN AnaVist, Version 2.0, now available with Derwent
MEMO	,	OHE	0 /	World Patents Index
NEWS	1.0	CED	13	FORIS renamed to SOFIS
NEWS		SEP		INPADOCDB enhanced with monthly SDI frequency
NEWS				CA/CAplus enhanced with monthly SDI frequency
NEWS	12	SEP	1 /	1967-1998
NEWS	13	SEP	17	CAplus coverage extended to include traditional medicine
				patents
NEWS	14	SEP	2.4	EMBASE, EMBAL, and LEMBASE reloaded with enhancements
NEWS		OCT		CA/CAplus enhanced with pre-1907 records from Chemisches
				Zentralblatt
NEWS	16	OCT	19	BEILSTEIN updated with new compounds
NEWS		NOV		Derwent Indian patent publication number format enhanced
NEWS		NOV		WPIX enhanced with XML display format
NEWS		NOV		ICSD reloaded with enhancements
NEWS		DEC		LINPADOCDB now available on STN
NEWS		DEC		BEILSTEIN pricing structure to change
NEWS		DEC		USPATOLD added to additional database clusters
NEWS		DEC		IMSDRUGCONF removed from database clusters and STN
NEWS		DEC		DGENE now includes more than 10 million sequences
NEWS		DEC		TOXCENTER enhanced with 2008 MeSH vocabulary in
HEND	20	DEC	1,	MEDLINE segment
NEWS	26	DEC	17	MEDLINE and LMEDLINE updated with 2008 MeSH vocabulary
NEWS		DEC		CA/CAplus enhanced with new custom IPC display formats
NEWS		DEC		STN Viewer enhanced with full-text patent content
ишию	20	DEC	1,	from USPATOLD
NEWS	29	JAN	02	STN pricing information for 2008 now available
NEWS	30	JAN	16	CAS patent coverage enhanced to include exemplified
				prophetic substances
NEWS	31	JAN	28	USPATFULL, USPAT2, and USPATOLD enhanced with new
				custom IPC display formats
NEWS	32	JAN	28	MARPAT searching enhanced
NEWS		JAN		USGENE now provides USPTO sequence data within 3 days
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NEWS 34 JAN 28 TOXCENTER enhanced with reloaded MEDLINE segment
NEWS 35 JAN 28 MEDLINE and LMEDLINE reloaded with enhancements
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NEWS 36 FEB 08 STN Express, Version 8.3, now available

NEWS EXPRESS FEBRUARY 08 CURRENT WINDOWS VERSION IS V8.3. AND CURRENT DISCOVER FILE IS DATED 24 JANUARY 2008

NEWS HOURS STN Operating Hours Plus Help Desk Availability

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TOTAL

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FILE 'HOME' ENTERED AT 21:34:44 ON 14 FEB 2008

=> file CAPLUS

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=> e losartan

E1 LOSARTA/BI E2 LOSARTAM/BI E3 5490 --> LOSARTAN/BI E4 1 LOSARTANK/BI E5 1 LOSARTANS/BI 1 LOSARTANT/BI E6

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         9548 POLYMORPHS
        14667 POLYMORPH
                (POLYMORPH OR POLYMORPHS)
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The following are valid formats:
ABS ---- GI and AB
ALL ----- BIB, AB, IND, RE
APPS ----- AI, PRAI
BIB ----- AN, plus Bibliographic Data and PI table (default)
CAN ----- List of CA abstract numbers without answer numbers
CBIB ----- AN, plus Compressed Bibliographic Data
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DALL ----- ALL, delimited (end of each field identified)
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FAM ----- AN, PI and PRAI in table, plus Patent Family data
FBIB ----- AN, BIB, plus Patent FAM
IND ----- Indexing data
IPC ----- International Patent Classifications
MAX ----- ALL, plus Patent FAM, RE
PATS ----- PI, SO
SAM ----- CC, SX, TI, ST, IT
SCAN ----- CC, SX, TI, ST, IT (random display, no answer numbers;
             SCAN must be entered on the same line as the DISPLAY.
             e.g., D SCAN or DISPLAY SCAN)
STD ---- BIB, CLASS
IABS ----- ABS, indented with text labels
IALL ----- ALL, indented with text labels
IBIB ----- BIB, indented with text labels
IMAX ----- MAX, indented with text labels
ISTD ----- STD, indented with text labels
OBIB ----- AN, plus Bibliographic Data (original)
OIBIB ----- OBIB, indented with text labels
SBIB ----- BIB, no citations
SIBIB ----- IBIB, no citations
HIT ---- Fields containing hit terms
HITIND ----- IC, ICA, ICI, NCL, CC and index field (ST and IT)
             containing hit terms
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HITRN ----- HIT RN and its text modification
HITSTR ----- HIT RN, its text modification, its CA index name, and
             its structure diagram
HITSEQ ----- HIT RN, its text modification, its CA index name, its
            structure diagram, plus NTE and SEQ fields
FHITSTR ---- First HIT RN, its text modification, its CA index name, and
             its structure diagram
FHITSEQ ---- First HIT RN, its text modification, its CA index name, its
             structure diagram, plus NTE and SEO fields
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KWIC ----- Hit term plus 20 words on either side OCC ----- Number of occurrence of hit term and field in which it occurs

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All of the formats (except for SAM, SCAN, HIT, HITIND, HITRN, HITSTR, FHITSTR, HITSEO, FHITSEO, KWIC, and OCC) may be used with DISPLAY ACC to view a specified Accession Number. ENTER DISPLAY FORMAT (BIB): ibib ab

ANSWER 1 OF 14 CAPLUS COPYRIGHT 2008 ACS on STN 2007:761240 CAPLUS

ACCESSION NUMBER: 147:166619

DOCUMENT NUMBER:

TITLE: Preparation of B-amino acid derivatives as

dipeptidyl peptidase-IV inhibitors INVENTOR(S): Sattigeri, Jitendra A.; Ahmed, Shahadat; Andappan,

Murugaiah M. S.; Sethi, Sachin; Sharma, Lalima; Pal, Chanchal Kumar; Kandalkar, Sachin Ramesh; Mahajan, Dipak C.; Kishore, Kaushal; Bhatia, Sumati; Gadhave, Anil G.; Bansal, Vinay S.; Davis, Joseph Alexanand

PATENT ASSIGNEE(S): Ranbaxy Laboratories Limited, India

SOURCE: PCT Int. Appl., 74pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

	ENT				KIND		DATE			APPLICATION NO.						DATE			
						-													
WO	2007	0775	80		A2		2007	0712		WO 2	006-	IB55	006		2	0061	221		
WO	2007	0775	08		A3		2007	1025											
	W:	ΑE,	AG,	AL,	AM,	AT,	AU,	AZ,	BA,	BB,	BG,	BR,	BW,	BY,	BZ,	CA,	CH,		
		CN,	co,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,		
		GE,	GH,	GM,	GT,	HN,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KM,	KN,		
		KP,	KR,	KZ,	LA,	LC,	LK,	LR,	LS,	LT,	LU,	LV,	LY,	MA,	MD,	MG,	MK,		
		MX,	MY,	MZ,	NA,	NG,	NI,	NO,	NZ,	OM,	PG,	PH,	PL,	PT,	RO,				
		RS,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	SM,	SV,	SY,	TJ,	TM,	TN,	TR,	TT,		
		TZ,	UA,	UG,	US,	UZ,	VC,	VN,	ZA,	ZM,	ZW								
	RW:	AT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,	EE,	ES,	FI,	FR,	GB,	GR,	HU,	IE,		
	IS, IT, LT,					LV,	MC,	NL,	PL,	PT,	RO,	SE,	SI,	SK,	TR,	BF,	BJ,		
	CF, CG, CI,					GA,	GN,	GQ,	GW,	ML,	MR,	NE,	SN,	TD,	TG,	BW,	GH,		
	GM, KE, LS					MZ,	NA,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	AM,	AZ,	BY,		
	KG, KZ, MD					TJ,	TM,	AP,	EA,	EP,	OA								
PRIORITY	IORITY APPLN. INFO.:										005-		A 20051230						
PRIORITY		GE, KP, MN, RS, TZ, AT, IS, CF, GM, KG,	GH, KR, MW, RU, UA, BE, IT, CG, KE, KZ,	GM, KZ, MX, SC, UG, BG, LT, CI, LS, MD,	GT, LA, MY, SD, US, CH, LU, CM, MW, RU,	HN, LC, MZ, SE, UZ, CY, LV, GA, MZ,	HR, LK, NA, SG, VC, CZ, MC, GN, NA,	HU, LR, NG, SK, VN, DE, NL, GQ, SD, AP,	ID, LS, NI, SL, ZA, DK, PL, GW, SL, EA,	IL, NO, SM, ZM, EE, PT, ML, SZ, EP,	IN, LU, NZ, SV, ZW ES, RO, MR, TZ, OA	IS, LV, OM, SY, FI, SE, NE, UG,	JP, LY, PG, TJ, FR, SI, SN, ZM,	KE, MA, PH, TM, GB, SK, TD, ZW,	KG, MD, PL, TN, GR, TR, TG, AM,	KM, MG, PT, TR, HU, BF, BW, AZ,	KN, MK, RO, TT, IE, BJ, GH, BY,		

OTHER SOURCE(S): MARPAT 147:166619

AB The invention relates to the preparation of  $\beta$ -amino acid derivs. I [A =

(hetero)aryl; E, E' = independently (CRaRb)n; n = 1-2; Ra, Rb = independently H, alk(en/yn)yl, cycloalkyl, (hetero)/aryl, heterocyclyl; RaCRb = optionally unsatd, ring; R = (un)substituted 2,5diazabicyclo[2.2.1]hept-2-yl, (piperidin-4-yl)amino, -3azabicyclo[3.1.0]hex-6-yl, etc.], and their pharmaceutically acceptable salts, pharmaceutically acceptable solvates, enantiomers, diastereomers, polymorphs, prodrugs, metabolites, and N-oxides, as dipeptidyl peptidase-IV inhibitors. This invention also relates to pharmacol. compns. containing the compds. of the invention, and methods of treating diabetes, especially type 2 diabetes, as well as prediabetes, diabetic dyslipidemia, metabolic acidosis, ketosis, satiety disorders, and obesity. These inhibitors can also be used to treat conditions manifested by a variety of metabolic, neurol., anti-inflammatory, and autoimmune disorders like inflammatory disease, multiple sclerosis, rheumatoid arthritis; viral, cancer and gastrointestinal disorders. I can also be used for treatment of infertility arising due to polycystic ovary syndrome. Thus, coupling 4-amino-1-[(morpholin-4-yl)carbonyl]piperidine tosylate with (3R)-3-[N-(tert-butoxycarbonyl)amino]-4-(2,4,5-trifluorophenyl)butanoic acid and cleavage of tert-butoxycarbonyl group in the presence of TFA gave II.TFA . I were evaluated for their peptidase-IV inhibitory activity using a fluorometric assay (IC50 values in the range of 1 nm to 10 µM).

## => d 12 2-8 ibib ab

L2 ANSWER 2 OF 14 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2006:1155444 CAPLUS

DOCUMENT NUMBER: 145:477884

TITLE: Angiotensin II receptor antagonists

INVENTOR(S): Alani, Laman L.; Dubost, David C.; Foster, Bruce S.;
Ghosh, Soumojeet; Jahansouz, Hossain; Pourkavoos,

Nazaneen; Rege, Bhagwant; Tatavarti, Aditya

PATENT ASSIGNEE(S): Merck & Co., Inc., USA

SOURCE: PCT Int. Appl., 34pp.

CODEN: PIXXD2
DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PA:	TENT	NO.			KIN	D	DATE			APPLICATION NO.						DATE			
WO	2006	1158	34		A1		2006	1102							2	0060	414		
	W:	AE,	AG,	AL,	AM,	AT,	AU,	AZ,	BA,	BB,	BG,	BR,	BW,	BY,	BZ,	CA,	CH,		
		CN,	co,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,		
		GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KM,	KN,	KP,	KR,		
		KZ,	LC,	LK,	LR,	LS,	LT,	LU,	LV,	LY,	MA,	MD,	MG,	MK,	MN,	MW,	MX,		
		MZ,	NA,	NG,	NI,	NO,	NZ,	OM,	PG,	PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,		
		SG,	SK,	SL,	SM,	SY,	ТJ,	TM,	TN,	TR,	TT,	TZ,	UA,	UG,	US,	UZ,	VC,		
		VN,	YU,	ZA,	ZM,	ZW													
	RW:	ΑT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,	EE,	ES,	FI,	FR,	GB,	GR,	HU,	IE,		
		IS,	IT,	LT,	LU,	LV,	MC,	NL,	PL,	PT,	RO,	SE,	SI,	SK,	TR,	BF,	BJ,		
		CF,	CG,	CI,	CM,	GA,	GN,	GQ,	GW,	ML,	MR,	ΝE,	SN,	TD,	TG,	BW,	GH,		
		GM,	KE,	LS,	MW,	MZ,	NA,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	AM,	ΑZ,	BY,		
		KG,	ΚZ,	MD,	RU,	ΤJ,	TM												
AU	2006	2402	47		A1		2006	1102		AU 2	006-	2402	47		2	0060	414		
									CA 2006-2604190							0060			
EP	1874	302			A1		2008	0109		EP 2	006-	7502	00		2	0060	414		
	R:	ΑT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,	EE,	ES,	FΙ,	FR,	GB,	GR,	HU,	IE,		
		IS,	IT,	LI,	LT,	LU,	LV,	MC,	NL,	PL,	PT,	RO,	SE,	SI,	SK,	TR			
IN	2007	CN04	215		A		2007	1221		IN 2	007-	CN42	15		2	0070	924		

US 2005-673086P P 20050420 WO 2006-US14092 W 20060414

The compds. of the present invention are polymorphic crystalline forms of the AB compound 2-butvl-4-chloro-1-[(2'-(1-H-tetrazol-5-vl)biphenvl-4yl)methyl]imidazole-5-carboxylic acid (I). Specifically, the compds. of the invention are selected from the group consisting of e.g., I, I-HCl Forms I through III and their monohydrate forms. I was prepared and converted to controlled release granules.

THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS REFERENCE COUNT: 2 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 3 OF 14 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2005:1004564 CAPLUS

DOCUMENT NUMBER: 143:292576

TITLE: Stabilization of a polymorphic form of

losartan potassium

INVENTOR(S): Svete, Peter; Grahek, Rok; Humar, Vlasta; Husu-Kovacevic, Breda; Jerala-Strukelj, Zdenka

PATENT ASSIGNEE(S): Lek Pharmaceuticals D.D., Slovenia PCT Int. Appl., 22 pp.

SOURCE: CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: Enalish

FAMILY ACC. NUM. COUNT: 2 PATENT INFORMATION:

P	PATENT		KIND DATE			APPLICATION NO.												
-						-									_			
W	TO 2005	0846	70		A1		2005	0915		WO 2	005-	EP21	80		2	0050	228	
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		CN,	CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,	
		GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KP,	KR,	KZ,	LC,	
		LK,	LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NA,	NI,	
		NO,	NZ,	OM,	PG,	PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	SM,	
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		AZ,	BY,	KG,	KZ,	MD,	RU,	TJ,	TM,	AT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,	
		EE,	ES,	FI,	FR,	GB,	GR,	HU,	ΙE,	IS,	IT,	LT,	LU,	MC,	NL,	PL,	PT,	
		RO,	SE,	SI,	SK,	TR,	BF,	ВJ,	CF,	CG,	CI,	CM,	GA,	GN,	GQ,	GW,	ML,	
		MR,	NE,	SN,	TD,	TG												
E	EP 1729	766			A1		2006	1213		EP 2	005-	7076	62		2	0050	228	
	R:	ΑT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,	EE,	ES,	FI,	FR,	GB,	GR,	HU,	ΙE,	
		IS,	IT,	LI,	LT,	LU,	MC,	NL,	PL,	PT,	RO,	SE,	SI,	SK,	TR			
Ü	JS 2007		A1 20071227					7 US 2007-590889						20070604				
PRIORI	. :						SI 2	004-	67			A 20040301						
										WO 2	0.05-	EP21	0.8	W 20050228				

Compns. were developed which stabilize an active pharmaceutical ingredient AB in polymorph form susceptible to degradation or interconversion into other polymorph forms, where stabilizing substance is conveniently among silicon dioxide, silicified microcryst. cellulose, magnesium oxide and polyethylene glycol. The polymorphic form of losartan potassium was stable when formulated with Syloid and PEG 6000.

REFERENCE COUNT: 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD, ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 4 OF 14 CAPLUS COPYRIGHT 2008 ACS on STN ACCESSION NUMBER: 2004:740317 CAPLUS

DOCUMENT NUMBER: 141:265973

TITLE: Preparation of polymorphic crystal forms of the antihypertensive agent losartan potassium

INVENTOR(S): Kumar, Pananchukunnath Manoj; Manikandan, Ramalingam; Singh, Romi Barat; Nagaprasad, Vishnubhotla; Malik,

Rajiv

PATENT ASSIGNEE(S): Ranbaxy Laboratories Limited, India

SOURCE: PCT Int. Appl., 22 pp. CODEN: PIXXD2

DOCUMENT TYPE: Patent English

LANGUAGE: FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

	ENT I				KIND DATE					APPL			DATE				
						_											
WO	2004	0764	42		A1		2004	0910	1	WO 2	004-	IB51	6		20	0040	227
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		BG,	BR,	BR,	BW,	BY,	BY,	BZ,	BZ,	CA,	CH,	CN,	CN,	CO,	CO,	CR,	CR,
		CU,	CU,	CZ,	CZ,	DE,	DE,	DK,	DK,	DM,	DZ,	EC,	EC,	EE,	EE,	EG,	ES,
		ES,	FI,	FI,	GB,	GD,	GE,	GE,	GH,	GM,	HR,	HR,	HU,	HU,	ID,	IL,	IN,
		IS,	JP,	JP,	KE,	KE,	KG,	KG,	KP,	KP,	KP,	KR,	KR,	KZ,	KZ,	KZ,	LC,
		LK,	LR,	LS,	LS,	LT,	LU,	LV,	MA,	MD,	MD,	MG,	MK,	MN,	MW,	MX,	MX,
		MZ,	MZ,	NA,	NI												
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		GQ,	GW,	ML,	, MR, NE, SN, TD,				TG								
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Pro	cess	es f	or p	rodu	lucing polymorphic of					stal	for	sarta	an				

potassium (I), useful as an antihypertensive, are claimed as are the crystal polymorphs of I, their crystal-characterization data,

and their use in pharmaceutical formulations.

REFERENCE COUNT: THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS 4 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 5 OF 14 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2004:740288 CAPLUS

DOCUMENT NUMBER: 141:248753

TITLE: Preparation of losartan potassium

polymorphs

INVENTOR(S): Boccignone, Andrea; Malpezzi, Luciana; Castaldi, Graziano; Allegrini, Pietro; Beltrame, Andrea

PATENT ASSIGNEE(S): Dinamite Dipharma S.P.A. In Abbreviate Form Dipharma

S.P.A., Italy; Dipharma S.P.A.

PCT Int. Appl., 25 pp. SOURCE:

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PA	TENT	NO.			KIND DATE					APPL	ICAT		DATE				
						_									-		
WO	2004	0764	06		A2		2004	0910		WO 2	004-	EP17	17		2	0040	220
WO	2004	0764	06		A3		2005	0113									
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		BG,	BR,	BR,	BW,	BY,	BY,	ΒZ,	ΒZ,	CA,	CH,	CN,	CN,	CO,	СО,	CR,	CR,
		CU,	CU,	CZ,	CZ,	DE,	DE,	DK,	DK,	DM,	DZ,	EC,	EC,	EE,	EE,	EG,	ES,
		ES,	FΙ,	FΙ,	GB,	GD,	GE,	GE,	GH,	GM,	HR,	HR,	HU,	HU,	ID,	IL,	IN,
		IS,	JP,	JP,	KE,	KE,	KG,	KG,	KP,	KP,	KP,	KR,	KR,	KZ,	KZ,	KZ,	LC,
	LK, LR, LS,					LT,	LU,	LV,	MA,	MD,	MD,	MG,	MK,	MN,	MW,	MX,	MX,
		MZ,	MZ,	NA,	NI												

RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG PRIORITY APPLN. INFO.: IT 2003-MI328 A 20030225 AB Losartan potassium polymorphs, identified as losartan potassium crystalline hydrate, losartan potassium amorphous and losartan potassium modification crystalline III, a process for their preparation, pharmaceutical compns, containing them and their use in therapy. Thus, losartan was dissolved in MeOh and treated with KHCO3 to give a losartan potassium polymorph III. ANSWER 6 OF 14 CAPLUS COPYRIGHT 2008 ACS on STN ACCESSION NUMBER: 2004:733484 CAPLUS DOCUMENT NUMBER: 142 - 127050 TITLE: Effect of enalapril and losartan on cytokines in patients with stable angina pectoris awaiting coronary artery bypass grafting and their interaction with polymorphisms in the interleukin-6 AUTHOR(S): Trevelvan, Jasper; Brull, David J.; Needham, Edward W. A.; Montgomery, Hugh E.; Morris, Alan; Mattu, Raj K. Department of Cardiology, University Hospitals of CORPORATE SOURCE: Coventry and Warwickshire, Coventry, UK American Journal of Cardiology (2004), 94(5), 564-569 SOURCE: CODEN: AJCDAG; ISSN: 0002-9149 PUBLISHER: Excerpta Medica, Inc. DOCUMENT TYPE: Journal LANGUAGE: English Angiotensin-converting enzyme inhibitors and angiotensin receptor blockers may have anti-inflammatory actions, an effect that could explain some of their beneficial effects on cardiovascular events in clin. trials. Coronary artery bypass grafting (CABG) is associated with a systemic inflammatory response and provides a convenient model to examine the effects of such agents. Genetic polymorphisms may be important in influencing the expression of cytokines, such as interleukin-6 (IL-6). We randomized men awaiting CABG to treatment with enalapril, losartan , or control for 2 mo before surgery. Systemic IL-6, IL-8, IL-10, and IL-1 receptor agonists were measured before and after surgery, and genotypes for the -174 G/C and -572 G/C IL-6 gene polymorphisms were determined Total release of the IL-1 receptor agonist was decreased 29% by enalapril and 31% by losartan (adjusted p = 0.041). IL-6 was decreased 17% by enalapril and 20% by losartan. Subjects possessing the -174 GG genotype produced 20% more IL-6 (adjusted p = 0.029). In these high producers of IL-6, release of IL-6 was decreased 51% by enalapril (adjusted p = 0.001) and 32% by losartan (adjusted p = 0.068).

has a similar but less marked effect. The production of IL-6 in this setting is influenced by the -174 G/C polymorphism.

REFERENCE COUNT: 19 THERE ARE 19 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

Release of IL-10 was nonsignificantly decreased 26% by enalapril and 21%

 $\rm IL-1$  receptor agonist after CABG. Enalapril produced a highly significant decrease of  $\rm 518$  in the release of  $\rm IL-6$  in patients identified as high producers of  $\rm IL-6$  by the -174 G/C polymorphism, whereas losartan

by losartan, whereas IL-8 was not detected. In conclusion, enalapril and losartan significantly decreased release of the

L2 ANSWER 7 OF 14 CAPLUS COPYRIGHT 2008 ACS on STN ACCESSION NUMBER: 2004:610104 CAPLUS

DOCUMENT NUMBER: 141:134092

TITLE: Telmisartan-simvastatin combination for the prophylaxis or treatment of cardiovascular,

cardiopulmonary, pulmonary, or renal diseases
INVENTOR(S): Riedel, Axel; Sendra, Josep-Maria; Leiter, Josef M.

E.; Kauschke, Stefan; Mark, Michael

PATENT ASSIGNEE(S): Boehringer Ingelheim International GmbH, Germany;

Boehringer Ingelheim Pharma GmbH & Co. Kg

SOURCE: PCT Int. Appl., 43 pp.
CODEN: PIXXD2

DOCUMENT TYPE: Patent
LANGUAGE: German

FAMILY ACC. NUM. COUNT: 6
PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. A1 20040729 WO 2004-EP175 \_\_\_\_\_\_ WO 2004062729 20040114 W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LK, LX, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MM, MX, MZ, NA
DE 10301372 A1 20040729 DE 2003-10301372 20030116
DE 10335027 A1 20050217 DE 2003-10335027 20030731
AU 2004204353 A1 20040729 AU 2004-204353 20040114
CA 2513281 A1 20040729 CA 2004-2513281 20040114
US 2004259925 A1 20041223 US 2004-757295 20040114
EP 1587584 B1 20070523 EP 2004-701918 20040114
EP 1587584 B1 20070523 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK BR 2004006812 A 20051227 BR 2004-6812 20040114 T 2006608 JP 2006-500558 AU 2004260606 A1 20050210 AU 2004-260606 CA 2534006 A1 20050210 CA 2004-2534006 EP 1651213 A1 20060503 EP 2004-763484 20040114 20040724 20040724 20040724 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, LB, S1, LI, LV, FI, RO, CY, TR, BG, CZ, EE, HU, PL, SK

CN 1829511

BR 2004013165

A 20060906

CN 2004-80022096

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JP 2007500677

T 20070118

JP 2006-521497

MX 2005PA07559

A 20050921

MX 2005PA07559

A 20050921

MX 2005PA0759

A 20050921

MX 2006PA01322

A 20060504

MX 2006-PA1322

A 20060504

MX 2006-PA1322

A 20060507

MX 20069A01322

A 20060227

RO 2006-938

A 20030116

DE 2003-10301372

A 2003016 DE 2003-10335027 A 20030731 DE 2003-1033027 A 20030731 DE 2003-10301371 A 20030116 US 2003-446695P P 20030211 US 2003-503317P P 20030216 DE 2003-10346260 A 20031006 DE 2003-10356815 A 20031205 WO 2004-EPR326 W 20040114 WO 2004-EP8326 W 20040724

AB The invention discloses a method for the prophylaxis or treatment of cardiovascular, cardiopulmonary, pulmonary or renal diseases, achieved by the improvement of endothelial function and the protection of organs, tissues and vessels when indications require a blood pressure check and a lipid level check, especially in patients that have been diagnosed with type 2 diabetes mellitus or if prediabetes is suspected. The method is also used for preventing diabetes and prediabetes and for the treatment of metabolic syndrome and insulin resistance in patients with normal blood pressure. The method involves the combined administration of effective quantities of telmisartan, or a polymorph or salt thereof, and simvastatin.

The invention also discloses suitable pharmaceutical compns. containing telmisartan, or a polymorph or salt thereof, and simvastatin, as a combined preparation for simultaneous, sep., or sequential use in the prophylaxis or treatment of the above diseases. Preparation of the sodium salt of telmisartan is described.

L2 ANSWER 8 OF 14 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2004:606351 CAPLUS

DOCUMENT NUMBER: 141:134089

TITLE: Telmisartan-atorvastatin combination for the prophylaxis or treatment of cardiovascular,

cardiopulmonary, pulmonary, or renal diseases
INVENTOR(S): Riedel, Axel; Sendra, Josep-Maria; Leiter, Josef M.

E.; Kauschke, Stefan; Mark, Michael

PATENT ASSIGNEE(S): Boohringer Ingelheim International GmbH, Germany; Boohringer Ingelheim Pharma GmbH & Co. Kg

SOURCE: PCT Int. Appl., 40 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: German

FAMILY ACC. NUM. COUNT: 6 PATENT INFORMATION:

	PAT	ENT	NO.			KIN	D	DATE			API	PLI	ICAT	ION	NO.		DATE				
	WO	2004	0625	57		A2	_	2004	0729		WO	20	004-	EP17	4		2	0040	114		
	WO	2004	0625	57		A3		2004	0916												
		W:						AU,													
			CN,	co,	CR,	CU,	CZ,	DK,	DM,	DZ,	, E0	С,	EE,	EG,	ES,	FI,	GB,	GD,	GE,		
								IL,											LK,		
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	DE	1030	1371			A1		2004	0805		DE	20	003-	1030	1371		2	20030116			
	DE	1033	5027			A1		2005	0217		DE	20	003-	1033	5027		20030731				
	ΑU	2004	2043	52		A1		2004	0729		AU	20	004-	2043	52		2	20040114			
	CA	2513	277			A1		2004	0729		CA	20	20040114								
	US	2004	2599	25		A1		2004	1223	AU 2004-204352 CA 2004-2513277 US 2004-757295 EP 2004-701904								20040114			
	EP	1587	479			A2		2005	1026		ΕP	20	004-	7019	04		2	0040	114		
		R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	, G1	R,	IT,	LI,	LU,	NL,	SE,	MC,	PT,		
	IE, SI				LT,	LV,	FI,	RO,	MK,	CY,	. Al	L,	TR,	BG,	CZ,	EE,	HU,	SK			
	BR	3R 2004006455				A		2005	1206		BR 2004-6455							0040	114		
	CN	1738	617		A		2006	0222	BR 2004-6455 CN 2004-80002407 JP 2006-500557 AU 2004-260606 CA 2004-2534006 EP 2004-763484								0040	114			
	JP	2006	5156	14		T	0601		JP	20	006-	5005	57		2	0040	114				
	AU	2004	2606	06		A1		AU	20	004-	2606	06		2	0040	724					
	CA	2534	006			A1		2005	0210	CA 2004-2534006							2	0040	724		
	EP	1651	213			A1		2006	0503		EP	20	004-	7634	84		2	0040	724		
		R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	, GI	R,	IT,	LI,	LU,	NL,	SE,	MC,	PT,		
			IE,	SI,	LT,	LV,	FI,	RO,	CY,	TR,	, B	3,	CZ,	EE,	HU,	PL,	SK				
	CN	1829	511			A		2006	0906		CN	20	004-	8002	2096		2	0040	724		
	BR	2004	0131	65		A		2006	1003		BR	2.0	004-	1316	5		- 2	0040	724		
	JP	2007	5006	77		T		2007	0118		JΡ	20	006-	5214	97		2	0040	724		
	$z_{A}$	2005	0035	42		A		2006	0726		ZA	20	005-	3542			2	0050	504		
	MV	2005	D307	102		70		2005	0026		MV	20	105	D 7 7 1	0.3		2	0050	620		
	IN 2005DN03073					A 20070112					E IN 2005-PN3073 6 NO 2005-3837 MX 2006-PA1322 7 NO 2006-938 DE 2003-10301371							20050711			
	NO 2005003837					A		2005	0815		NO	20	005-	3837			2	0050	815		
	MX 2006PA01322					A		2006	0504		MX	20	006-	PA13	22		2	0060	131		
	NO 2006000938					A		2006	0227		NO	20	006-	938			2	0060	227		
PRIOR	RIORITY APPLN. INF										DE	20	003-	1030	1371		A 2	0030	116		
											DE	20	003-	1033	5027		A 2	0030	731		

US 2003-446695P P 20030211 US 2003-503317P P 20030916 DE 2003-10346260 A 20031006 DE 2003-10356815 A 20031205 WO 2004-EP174 W 20040114 WO 2004-EP8326 W 20040724

The invention discloses a method for the prophylaxis or treatment of AB cardiovascular, cardiopulmonary, pulmonary, or renal diseases, achieved by the improvement of endothelial function and the protection of organs, tissues and vessels when indications require a blood pressure check and a lipid level check, especially in patients that have been diagnosed with type 2 diabetes mellitus or if prediabetes is suspected. The method is also used for preventing diabetes and prediabetes and for the treatment of metabolic syndrome and insulin resistance in patients with normal blood pressure. The method involves the combined administration of effective amts. of telmisartan, or a polymorph or salt thereof, and atorvastatin. The invention also discloses suitable pharmaceutical compns. containing telmisartan, or a polymorph or salt thereof, and atorvastatin, as a combined preparation for simultaneous, sep. or sequential use in the prophylaxis or treatment of the above diseases. Preparation of the sodium salt of telmisartan is described.

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COST IN U.S. DOLLARS SINCE FILE TOTAL SESSION ENTRY FULL ESTIMATED COST 29.92 30.13 DISCOUNT AMOUNTS (FOR OUALIFYING ACCOUNTS) SINCE FILE TOTAL ENTRY SESSION CA SUBSCRIBER PRICE -6.40 -6.40

FILE 'STNGUIDE' ENTERED AT 21:37:53 ON 14 FEB 2008 USE IS SUBJECT TO THE TERMS OF YOUR CUSTOMER AGREEMENT COPYRIGHT (C) 2008 AMERICAN CHEMICAL SOCIETY (ACS)

FILE CONTAINS CURRENT INFORMATION. LAST RELOADED: Feb 8, 2008 (20080208/UP).

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The search profile that was entered contains terms or nested terms that are not separated by a logical operator.

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L2 ANSWER 9 OF 14 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2004:414643 CAPLUS

DOCUMENT NUMBER: 140:412339

TITLE: Crystalline form of losartan potassium

Reddy, Manne Satyanarayana; Eswaraiah, Sajja; Koppera, INVENTOR(S):

Ravinder Reddy; Reddy, Vajrala Venkata PATENT ASSIGNEE(S): Reddy's Laboratories Limited, India; Reddy's

Laboratories, Inc. SOURCE:

U.S. Pat. Appl. Publ., 11 pp. CODEN: USXXCO

DOCUMENT TYPE: Patent LANGUAGE: English FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE A1 20040520 US 2003-629316 20030729 a 20070727 IN 2002-MA568 20020729 US 2004097568 IN 2002MA00568 20070727 IN 2002-MA568 20020729 IN 2002-MA568 A 20020729 PRIORITY APPLN. INFO.:

AB A compound that is a crystalline Form III of losartan potassium is provided. Also provided are compns. containing the compound and methods for

its preparation For example, 125 g of trityl losartan (preparation given) was

mixed with an aqueous solution containing 11 g of KOH, 125 mL water, and 1250 mT.

methanol until the reaction was complete. The solvent was distilled off the reaction solution under vacuum, and water (325 mL) added to the residual mass, stirred for 30 min, the pH adjusted to 8.2 to 8.8, and the mass filtered. The filtrate was washed with water, the water was distilled off, and the resulting residue was dissolved in methanol, the solvent distilled off, and the residual mass cooled to a temperature of 5 to 10°, filtered, and dried to yield crystalline polymorph Form III of losartan potassium (weight 43.0 g). The crystalline polymorph Form III of losartan potassium was also obtained from crystalline polymorph Form I of losartan potassium.

ANSWER 10 OF 14 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2001:287433 CAPLUS

DOCUMENT NUMBER: 136:58666

Polymorphic transformation of losartan TITLE:

AUTHOR(S): Elbary, A. Abd; Nafadi, M. M.; El-Khateeb, Mona A. Department of pharmaceutics, Faculty of Pharmacy, CORPORATE SOURCE:

Cairo University, Cairo, Egypt

Egyptian Journal of Pharmaceutical Sciences (2000), SOURCE: Volume Date 1999, 40(1), 49-59

CODEN: EJPSBZ; ISSN: 0301-5068

PUBLISHER: National Information and Documentation Centre

DOCUMENT TYPE: Journal

LANGUAGE: English

AB Losartan was recrystd. from alc. solns. containing each of PEG 4000, PEG 6000, Tween 40, Tween 80, Myri 59 and PVPK90 sep. Samples of the drug, with or without polymers or surfactants were investigated using DSC, XRD, IR and microphotographs to reveal the presence or absence of polymorphism. There were polymorphic changes of the drug with all the tested additives and all polymorphic forms are crystalline

THERE ARE 20 CITED REFERENCES AVAILABLE FOR THIS REFERENCE COUNT: 20 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 11 OF 14 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 1997:90112 CAPLUS DOCUMENT NUMBER: 126:297598

SOLUBILITY SOLUBILITY

CODEN: PHARAT; ISSN: 0031-7144 PUBLISHER: Govi-Verlag Pharmazeutischer Verlag

DOCUMENT TYPE: Journal LANGUAGE: English

AB The solubilities of the enantiotropic polymorphs I and II of losartan, an orally administered angiotensin II receptor

antagonist used in the treatment of hypertension, was determined to evaluate the free energy differences between the two forms. The solubilities of form I and II at 25-65° were 0.59-1.25 and 1.95-2.63 resp. Thus, form I has lower solubility and is more thermodynamically stable.

L2 ANSWER 12 OF 14 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 1995:890142 CAPLUS

DOCUMENT NUMBER: 123:313978

TITLE: Polymorphs of losartan potassium

and a process for the preparation of polymorph forms I and II of losartan potassium

INVENTOR(S): Campbell, Gordon Creston, Jr.; Dwivedi, Anil M.;

Levorse, Dorothy A.; McCauley, James A.; Raghavan,

Krishnaswamy S.

PATENT ASSIGNEE(S): Merck and Co., Inc., USA; du Pont de Nemours, E. I.,

and Co.; Dupont Merck Pharmaceutical Co. Patent.

English

PCT Int. Appl., 54 pp. SOURCE: CODEN: PIXXD2

DOCUMENT TYPE: LANGUAGE:

FAMILY ACC. NUM. COUNT: 1 PATENT INFORMATION:

E	PA:	ENT:	NO.			KIND DATE					APPL	ICAT		DATE						
Ţ-	10	9517	396			A1	_	1995	0629		WO 1	994-	US14	768		1	9941	221		
		W:	AM,	AU,	BB,	BG,	BR,	BY,	CA,	CN,	CZ,	EE,	FI,	GE,	HU,	JP,	KG,	KR,		
			KZ,	LK,	LR,	LT,	LV,	MD,	MG,	MN,	NO,	NZ,	PL,	RO,	RU,	SI,	SK,	TJ,		
			TT,	UA,	US,	UZ														
		RW:	KE,	MW,	SD,	SZ,	ΑT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	ΙE,	IT,	LU,		
			MC,	NL,	PT,	SE,	BF,	ВJ,	CF,	CG,	CI,	CM,	GA,	GN,	ML,	MR,	NE,	SN,		
			TD,	TG																
(	ĊΑ	2179	067			A1		1995	0629		CA 1	994-	2179	067		1	9941	221		
P	U	9514	058			A		1995	0710		AU 1	995-	1405	8		1	9941	221		
P	U	6858	98			B2		1998	0129											
E	EΡ	7360	21			A1		19961009			EP 1995-905		9054	5449		1	9941	221		
	R: AT, BE, CH							ES,	FR,	GB,	GR,	IE,	IT,	LI,	LU,	NL,	PT,	SE		
ن	JP 09507075							1997	0715	JP 1994-517594						19941221				
Ţ	5608	075			A 19970304				US 1995-371937						19950112					
PRIORI	PRIORITY APPLN. INFO.:											US 1993-173440						223		
										WO 1994-US14768						W 19941221				

Polymorphic forms of losartan potassium, I, a known angiotensin II-inhibiting antihypertensive, are prepared Numerous spectral, thermal, and X-ray data of I form I and II are reported, and I-containing formulations are presented along with angiotensin II receptor inhibition data.

L2 ANSWER 13 OF 14 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 1994:116612 CAPLUS

DOCUMENT NUMBER: 120:116612

TITLE: Thermal analysis and solution calorimetry studies on losartan polymorphs

AUTHOR(S): Wu, Lei Shu; Gerard, Christine; Hussain, Munir A. CORPORATE SOURCE: Du Pont Merck Pharm, Co., Wilmington, DE, 19880-0336,

USA

Pharmaceutical Research (1993), 10(12), 1793-5 SOURCE: CODEN: PHREEB: ISSN: 0724-8741

DOCUMENT TYPE: Journal

LANGUAGE: English The existence of losartan (I) polymorphs was confirmed

by solution calorimetry. The heat of transition was determined to be 1.74 kcal/mol from Form I to Form II. Form I is thermodynamically more stable

than Form II at ambient temperature Form II could convert to Form I during storage at ambient temperature since the conversion is exothermic.

L2 ANSWER 14 OF 14 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 1993:525045 CAPLUS

DOCUMENT NUMBER: 119:125045

TITLE: A spectroscopic investigation of Losartan

polymorphs

AUTHOR(S): Raghavan, Krishnaswamy; Dwivedi, Anil; Campbell, G.

Creston, Jr.; Johnston, Eric; Levorse, Dorothy;

McCauley, James; Hussain, Munir

CORPORATE SOURCE: Exp. Stn., Du Pont Merck Pharm. Co., Wilmington, DE,

19880-0400, USA

SOURCE: Pharmaceutical Research (1993), 10(6), 900-4

CODEN: PHREEB; ISSN: 0724-8741

DOCUMENT TYPE: Journal LANGUAGE .

English

Losartan (I), an antihypertensive agent in clin. development,

existed in 2 enantiotropic polymorphic forms, a low-temperature stable form (Form I) and a high-temperature stable form (Form II), the temps. at which they are stable being related to the transition temperature X-ray powder

diffraction

patterns indicated differences in the crystal packing of the 2 forms. The vibrational data from IR and Raman spectroscopy suggested a subtle change in mol. conformation and crystal packing in the 2 forms. Solid-state 13C NMR data of the polymorphs concurred with the vibrational data and indicated that, while the observed line widths reflect no major changes in crystallinity, signal multiplicities and chemical shifts do reflect differences in mol. packing in the resp. unit cells. Thus, in the absence of crystallog. data, useful structural information could be derived from

spectroscopic results to identify each of the crystalline forms.